



RAW SEQUENCE LISTING

PATENT APPLICATION: US/09/554,414B

DATE: 11/12/2001 TIME: 18:23:51

```
4 <110> APPLICANT: McGILL UNIVERSITY
      5
              SZYF, Moshe
      6
              BHATTACHARYA, Sanjoy K.
      7
              RAMCHANDANI, Shyam
     10 <120> TITLE OF INVENTION: DNA DEMETHYLASE, THERAPEUTIC AND
     11
              DIAGNOSTIC USES THEREOF
     13 <130> FILE REFERENCE: 1770-183"PCT" FC/ld
C--> 15 <140> CURRENT APPLICATION NUMBER: US/09/554,414B
C--> 15 <141> CURRENT FILING DATE: 2000-09-06
     15 <150> PRIOR APPLICATION NUMBER: CA 2,220,805
     16 <151> PRIOR FILING DATE: 1997-11-12
                                                                 ENTERED
     18 <150> PRIOR APPLICATION NUMBER: CA 2,230,991
     19 <151> PRIOR FILING DATE: 1998-05-11
     21 <160> NUMBER OF SEQ ID NOS: 10
     23 <170> SOFTWARE: FastSEQ for Windows Version 3.0
    25 <210> SEQ ID NO: 1
    26 <211> LENGTH: 1804
    27 <212> TYPE: DNA
    28 <213> ORGANISM: Unknown
    30 <220> FEATURE:
    31 <223> OTHER INFORMATION: cDNA encoding human demethylase
    33 <400> SEQUENCE: 1
    34 ccgctctgcg ggcggggcgg gtctccggga ttccaagggc tcggttacgg aagaagcgca
                                                                                60
    35 gagccggctg gggaggggc tggatgcgcg cgcacccggg gggaggccgc tgctgcccgg
                                                                               120
    36 agcaggagga gggggagagc gcggcggggg gcagcggcgc tggcggcgac tccgccatag
                                                                               180
    37 agcagggggg ccagggcagc gcgctcgctc cgtccccggt gagcggcgtg cgcagggaag
                                                                               240
    38
        gcgctcgggg cggcggccgt ggccgggggc ggtggaagca ggcggcccgg ggcggcggcg
                                                                               300
        tetgtggccg tggccgtggc cgtggccggg gtcggggccg tggccggggc cggggccggg
    39
                                                                               360
    40 gccgcggccg tccccagagt ggcggcagcg gccttggcgg cgacggcggc ggcggcgcgg
                                                                              420
    41 gcggctgcgg cgtcggcagc ggtggcggcg tcgccccccg gcgggatcct gtccctttcc
                                                                              480
    42 cgtcggggag ctcggggccg gggcccaggg gaccccgggc cacggagagc gggaagagga
                                                                              540
    43 tggactgccc ggccctcccc cccggatgga agaaggagga agtgatccga aaatcagggc
                                                                              600
    44 tcagtgctgg caagagcgat gtctactact tcagtccaag tggtaagaag ttcagaagta
                                                                              660
    45 aacctcagct ggcaagatac ctgggaaatg ctgttgacct tagcagtttt gacttcagga
                                                                              720
    46
        ccggcaagat gatgcctagt aaattacaga agaacaagca gagactccgg aatgaccccc
                                                                              780
        tcaatcagaa caagggtaaa ccagacctga acacaacatt gccaattaga caaactgcat
    47
                                                                              840
        caattttcaa gcaaccagta accaaattca cgaaccaccc gagcaataag gtgaagtcag
    48
                                                                              900
        acccccagcg gatgaatgaa caaccacgtc agcttttctg ggagaagagg ctacaaggac
    49
                                                                              960
    50
       ttagcgcatc agatgtaaca gaacaaatta taaaaaccat ggagctacct aaaggtcttc
                                                                             1020
    51
        aaggagtegg teeaggtage aatgacgaga eeettetgte tgetgtggee agtgetttae
                                                                             1080
    52 acacaagete tgegeceate acaggacaag tetetgetge egtggaaaag aaccetgetg
                                                                             1140
    53 tttggcttaa cacatctcaa cccctctgca aagctttcat tgttacagat gaagacatta
                                                                             1200
        ggaaacagga agagcgagtc caacaagtac gcaagaaact ggaggaggca ctgatggccg
                                                                             1260
    55
        acatectgte eegggetgeg gacaeggagg aagtagacat tgacatggae agtggagatg
                                                                             1320
       aggcgtaaga atatgatcag gtaactttcg actgaccttc cccaagagca aattgctaga
                                                                             1380
        aacagaatta aaacatttcc actgggtttc gcctgtaaga aaaagtgtac ctgagcacat
                                                                             1440
    58 agctttttaa tagcactaac caatgccttt ttagatgtat ttttgatgta tatatctatt
                                                                             1500
```

RAW SEQUENCE LISTING DATE: 11/12/2001 PATENT APPLICATION: US/09/554,414B TIME: 18:23:51

59 60 61 62 63	cagggccett ceggtgeagt geagetttga ggceaggtge agtetactgg aaaggtagea ] cttacgtgaa atatttgttt ceceeacagt tttaatataa acagateagg agtaceaaat ] aagttteeca attaaagatt attataette actgtatata aacagatttt tataetttat ]														1620 1680 1740		
64	tcac				c y cu	Ju C.		ccac	c ac	cacc	y caa	aga	caaa	Laa	acyai	Juacac	1804
66	<210	> SE	Q ID	NO:	2												1004
	<211																
68	<212	> TY	PE:	PRT													
	<213				Jnkno	own											
	<220>																
						ON:	pre	dict	ed ai	nino	acio	d of	huma	an d	emetl	nylase	
	<400>																
75		Arg	Ala	His	_	Gly	Gly	Gly	Arg		Cys	Pro	Glu	Gln	Glu	Glu	
76	1	<b>01</b>	<b>a</b>		5			_		10					15	_	
77 70	СТА	GIU	ser		АТа	GLY	GLY	Ser		Ala	Gly	GLY	Asp		Ala	Ile	
78 79	Clu	Cln	C1	20	C1 n	<b>~1</b>	<b>G</b>	31 <u>-</u>	25	<b>3</b> 3 -	n	<b>~</b>		30	~		
80	GIU	GIII	35	СТА	GIII	GIY	ser		ьeu	Ата	Pro	ser		vaı	Ser	GLY	
81	Va l	Δra		Glu.	Glw	λla	λνα	40	C1,,	C1**	7 mm	C1	45	C1	Arg	(T)	
82	VUL	50	пту	GIU	GIY	АІа	55	СТУ	GLY	СТУ	AIG	60	Arg	GTÀ	Arg	тгр	
83	Lvs		Ala	Glv	Ara	Glv		Glv	Val	Cve	Glv		G1v	λνα	Gly	λκα	
84	65			1	9	70	<b>0</b> -1	<b>U 1</b>	, 41	CIS	75	nrg	GLY	. Tu	GIY	80	
85	Gly	Arq	Gly	Arq	Gly	Arq	Gly	Arq	Glv	Ara		Ara	Glv	Ara	Gly		
86	-		_	,	85	,	1		1	90	~- <i>1</i>	5		9	95	9	
87	Pro	Pro	Ser	Gly	Gly	Ser	Gly	Leu	Gly	Gly	Asp	Gly	Gly	Gly	Cys	Glv	
88				100	_		-		105	•	•	4	-	110	- 2 -	1	
89	Gly	Gly	Gly	Ser	Gly	Gly	Gly	Gly	Ala	Pro	Arg	Arg	Glu	Pro	Val	Pro	
90			115					120					125				
91	Phe		Ser	Gly	Ser	Ala	Gly	Pro	Gly	Pro	Arg	Gly	Pro	Arg	Ala	Thr	
92		130	<b>-</b>				135					140					
93		Ser	GLy	Lys	Arg		Asp	Cys	Pro	Ala		Pro	Pro	Gly	Trp	_	
94	145	<b>a</b> 1	<b>01</b>	** - 1		150	_	_		_	155					160	
95 96	Lys	GIU	GIU	vaı		Arg	Lys	ser	GTĀ		Ser	Ala	GLy	Lys	Ser	Asp	
90 97	Val	Фил	Пттх	Dho	165	Dro	Co~	<i>c</i> 1	T	170	nh -	3	<b>a</b>	<b>.</b>	175	<b>63</b>	
98	Val	тут	ıyı	180	261	PIO	ser	СТУ	LуS 185	гàг	Pne	Arg	ser	Lуs 190	Pro	GIn	
99	Leu	Ala	Ara		Leu	Glv	Δen	Thr		λen	Lau	cor	Sor		Asp	Dho	
100	204		195		ьси	011	non	200		пор	пец	SET	205		ASP	PHE	
101	Arg	Thr			Met	Met	Pro			Len	Gln	Lvs			: Gln	Arg	
102	,	210		-1-			215		-70	200	. 0111	220		. Dy c	, 011	nrg	
103	Leu	Arg	Asn	Asp	Pro	Leu			Asn	Lys	Gly			Asp	Leu	Asn	
104	225			_		230				•	235					240	
105	Thr	Thr	Leu	Pro	Ile	Arg	Gln	Thr	Ala	Ser	Ile	Phe	Lys	Gln	Pro	Val	
106					245					250			-		255		
107	Thr	Lys	Val			His	Pro	Ser	Asn	Lys	Val	Lys	Ser	Asp	Pro	Gln	
108				260					265					270			
109	Arg	Met			Gln	Pro	Arg			Phe	Trp	Glu	Lys	Arg	Leu	Gln	
110			275					280					285				

RAW SEQUENCE LISTING

DATE: 11/12/2001 PATENT APPLICATION: US/09/554,414B TIME: 18:23:51

```
111
      Gly Leu Ser Ala Ser Asp Val Thr Glu Gln Ile Ile Lys Thr Met Glu
 112
          290
                               295
                                                   300
      Leu Pro Lys Gly Leu Gln Gly Val Gly Pro Gly Ser Asn Asp Glu Thr
 113
 114
      305
                           310
                                               315
                                                                   320
      Leu Leu Ser Ala Val Ala Ser Ala Leu His Thr Ser Ser Ala Pro Ile
 115
 116
                      325
                                           330
      Thr Gly Gln Val Ser Ala Ala Val Glu Lys Asn Pro Ala Val Trp Leu
 117
 118
                  340
                                       345
      Asn Thr Ser Gln Pro Leu Cys Lys Ala Phe Ile Val Thr Asp Glu Asp
 119
 120
                                   360
                                                       365
 121
      Ile Arg Lys Gln Glu Glu Arg Val Gln Gln Val Arg Lys Leu Glu
 122
          370
                               375
                                                   380
 123
      Glu Ala Leu Met Ala Asp Ile Leu Ser Arg Ala Ala Asp Thr Glu Glu
 124
                          390
                                               395
 125
      Met Asp Ile Glu Met Asp Ser Gly Asp Glu Ala
 126
                      405
 128 <210> SEQ ID NO: 3
 129 <211> LENGTH: 1589
 130 <212> TYPE: DNA
131 <213> ORGANISM: Unknown
133 <220> FEATURE:
134 <223> OTHER INFORMATION: cDNA sequence of human dMTase2
136 <400> SEQUENCE: 3
137
     cacgcgcggg cgggtgggcg gagcggcccc cctagcgggg gctgtgaagc gcggggaggg
                                                                              60
     ggccgagcgg gtggcgaagc cggcgcgcg ccggctgggg gcggagggcg gaggcccgtg
138
                                                                             120
139
     ggacagaaca gctgcggcga gtggcggcgg cggagggagc cgaatcggcg acgagcccgg
                                                                             180
140
     gggtcgcaac ttgcagaagc ggcggcggcg gcggcatcgg ccacggcggg cggaaaagcc
                                                                             240
141
     ggggcgcaat ggagcggaag aggtgggagt gcccggcgct cccgcagggc tgggaaaggg
                                                                             300
142
     aagaagtgcc caggaggtcg gggctgtcgg ccggccacag ggatgtcttt tactatagcc
                                                                             360
     ccagcgggaa gaagttccgc agcaagccac aactggcacg ttacctgggc ggatccatgg
143
                                                                             420
144
     acctcagcac cttcgacttc cgcaccggaa agatgttgat gaacaagatg aataagagtc
                                                                             480
145
     gccagcgtgt gcgctatgat tettecaace aggteaaggg caageetgae etgaacaeeg
                                                                             540
146
     cgctgcctgt acggcagact gcatccatct tcaagcaacc ggtgaccaag atcaccaacc
                                                                             600
147
     accccagcaa caaggtcaag agcgacccgc agaaggcagt ggaccagccg aggcagcttt
                                                                             660
148
     tctgggagaa gaagctaagt ggattgagtg cctttgacat tgcagaagaa ctggtcagga
                                                                             720
149
     ccatggactt gcccaagggc ctgcagggag tgggccctgg ctgtacagat gagacgctgc
                                                                             780
     tgtcagccat tgcgagtgct ctacacacca gcaccctgcc cattacaggc cagctctctg
150
                                                                             840
151
     cageegtgga gaagaaceet ggtgtgtgge tgaacaetge acageeactg tgcaaageet
                                                                             900
152
     tcatggtgac agatgacgac atcaggaagc aggaggagct ggtacagcag gtacggaagc
                                                                            960
     gcctggagga ggcactgatg gccgacatgc tagctcatgt ggaggagctt gcccgagacg
153
                                                                           1020
     gggaggcacc actggacaag gcctgtgcag aggaggaaga ggaggaggaa gaggaggagg
                                                                           1080
155
     aagagccgga gccagagcga gtgtagcaca ggtgccctgc ccaagtctgg gctgcagact
                                                                           1140
     gccttcagcc ttgcctggac caggtagggg ccagacctgt aggaggcagc cgtccacctc
156
                                                                           1200
     ctttccaaag cctcctgctt ccaggtctca gtgcagggag cccctgtgga ccttgaactc
157
                                                                           1260
    acttgtccct gcgctgcctg gcaggaagcc ccacactgaa agcagatgag cagtgaccca
158
                                                                           1320
159
     actgagagge cacctggaca cagteacete cetgeeteet tateatagga caaggeettg
                                                                           1380
160
     cttggcaccg aggagetggg ageegtgttg ggtgetggag gaagtttetg gaaacacac
                                                                           1440
    tggctatgcc caccttatgt ccctaaggct attacaggcc agggtttgga ctgctccggc
161
                                                                           1500
    ccacaggget gcccagcete cccacactga gggtcagcag cccaccagga agtcacttte
162
                                                                           1560
```

RAW SEQUENCE LISTING DATE: 11/12/2001 PATENT APPLICATION: US/09/554,414B TIME: 18:23:51

```
163 cttcaataaa ctgatggtag gaacttgtg
                                                                             1589
 165 <210> SEQ ID NO: 4
 166 <211> LENGTH: 291
 167 <212> TYPE: PRT
 168 <213> ORGANISM: Unknown
 170 <220> FEATURE:
 171 <223> OTHER INFORMATION: predicted amino acid sequence of human dMTase2
 173 <400> SEQUENCE: 4
      Met Glu Arg Lys Arg Trp Glu Cys Pro Ala Leu Pro Gln Gly Trp Glu
 175
 176
      Arg Glu Glu Val Pro Arg Arg Ser Gly Leu Ser Ala Gly His Arg Asp
 177
                  20
                                      25
 178
     Val Phe Tyr Tyr Ser Pro Ser Gly Lys Lys Phe Arg Ser Lys Pro Gln
179
     Leu Ala Arg Tyr Leu Gly Gly Ser Met Asp Leu Ser Thr Phe Asp Phe
180
181
182
     Arg Thr Gly Lys Met Leu Met Ser Lys Met Asn Lys Ser Arg Gln Arg
183
                                              75
     Val Arg Tyr Asp Ser Ser Asn Gln Val Lys Gly Lys Pro Asp Leu Asn
184
185
186
     Thr Ala Leu Pro Val Arg Gln Thr Ala Ser Ile Phe Lys Gln Pro Val
187
                                      105
     Thr Lys Ile Thr Asn His Pro Ser Asn Lys Val Lys Ser Asp Pro Gln
188
189
                                  120
190
     Lys Ala Val Asp Gln Pro Arg Gln Leu Phe Trp Glu Lys Lys Leu Ser
191
                              135
                                                  140
192
     Gly Leu Asn Ala Phe Asp Ile Ala Glu Glu Leu Val Lys Thr Met Asp
193
                         150
                                              155
     Leu Pro Lys Gly Leu Gln Gly Val Gly Pro Gly Cys Thr Asp Glu Thr
194
195
                                          170
196
     Leu Leu Ser Ala Ile Ala Ser Ala Leu His Thr Ser Thr Met Pro Ile
197
     Thr Gly Gln Leu Ser Ala Ala Val Glu Lys Asn Pro Gly Val Trp Leu
198
199
                                                      205
200
     Asn Thr Thr Gln Pro Leu Cys Lys Ala Phe Met Val Thr Asp Glu Asp
201
                             215
202
     Ile Arg Lys Gln Glu Glu Leu Val Gln Gln Val Arg Lys Arg Leu Glu
203
                         230
                                              235
204
     Glu Ala Leu Met Ala Asp Met Leu Ala His Val Glu Glu Leu Ala Arg
205
                     245
                                          250
206
     Asp Gly Glu Ala Pro Leu Asp Lys Ala Cys Ala Glu Asp Asp Asp Glu
207
                 260
                                     265
208
     Glu Asp Glu Glu Glu Glu Glu Glu Pro Asp Pro Asp Pro Glu Met
209
             275
                                 280
210
    Glu His Val
211
         290
213 <210> SEQ ID NO: 5
214 <211> LENGTH: 1966
215 <212> TYPE: DNA
```

RAW SEQUENCE LISTING DATE: 11/12/2001 PATENT APPLICATION: US/09/554,414B TIME: 18:23:51

```
216 <213> ORGANISM: Unknown
218 <220> FEATURE:
219 <223> OTHER INFORMATION: cDNA sequence of mouse dMTasel
221 <400> SEQUENCE: 5
     gggggcgtgg ccccgagaag gcggagacaa gatggccgcc catagcgctt ggaggaccta
                                                                             60
     agaggcggtg gccggggcca cgccccgggc aggagggccg ctctgtgcgc gcccgctcta
223
                                                                            120
224
     tgatgcttgc gcgcgtcccc cgcgcgccgc gctgcgggcg gggcgggtct ccgggattcc
                                                                            180
225
     aagggctcgg ttacggaaga agcgcagcgc cggctgggga gggggctgga tgcgcgcgca
                                                                            240
226
     cccgggggga ggccgctgct gcccggagca ggaggggg gagagtgcgg cgggcggcag
                                                                            300
227
     eggegetgge ggegaeteeg ceatagagea ggggggeeag ggeagegege tegeeeegte
                                                                            360
     cccggtgagc ggcgtgcgca gggaaggcgc tcggggcggc ggccgtggcc gggggcggtg
                                                                            420
229
     gaagcaggcg ggccggggcg gcggcgtctg tggccgtggc cggggccggg gccgtggccg
                                                                            480
230
     gggacgggga cggggccggg gccggggccg cggccgtccc ccgagtggcg gcagcggcct
                                                                            540
231
     tggcggcgac ggcggcggct gcggcggcgg cggcagcggt ggcggcggcg ccccccggcg
                                                                            600
232
     ggagccggtc cctttcccgt cggggagcgc ggggccgggg cccaggggac cccgggccac
                                                                            660
233
     ggagagcggg aagaggatgg attgcccggc cctcccccc ggatggaaga aggaggaagt
                                                                            720
234
     gatccgaaaa tctgggctaa gtgctggcaa gagcgatgtc tactacttca gtccaagtgg
                                                                            780
235 taagaagttc agaagcaagc ctcagttggc aaggtacctg ggaaatactg ttgatctcag
                                                                            840
236 cagttttgac ttcagaactg gaaagatgat gcctagtaaa ttacagaaga acaaacagag
                                                                            900
237
     actgcgaaac gatcctctca atcaaaataa gggtaaacca gacttgaata caacattqcc
                                                                            960
238 aattagacaa acagcatcaa ttttcaaaca accggtaacc aaagtcacaa atcatcctag
                                                                           1020
239
     taataaagtg aaatcagacc cacaacgaat gaatgaacag ccacqtcagc ttttctqqqa
                                                                           1080
240
     gaagaggcta caaggactta gtgcatcaga tgtaacagaa caaattataa aaaccatgga
                                                                           1140
241
     actacccaaa ggtcttcaag gagttggtcc aggtagcaat gatgagaccc ttttatctgc
                                                                           1200
242 tgttgccagt gctttgcaca caagctctgc gccaatcaca gggcaagtct ccgctgctgt
                                                                           1260
243
     ggaaaagaac cctgctgttt ggcttaacac atctcaaccc ctctgcaaag cttttattgt
                                                                           1320
244 cacagatgaa gacatcagga aacaggaaga gcgagtacag caagtacgca agaaattgga
                                                                           1380
     agaagcactg atggcagaca tcttgtcgcg agctgctgat acagaagaga tggatattga
                                                                           1440
246
     aatggacagt ggagatgaag cctaagaata tgatcaggta actttcgacc gactttcccc
                                                                           1500
     aagrgaaaat tootagaaat tgaacaaaaa tgtttocact ggottttgcc tgtaagaaaa
247
                                                                           1560
     aaaatgtacc cgagcacata gagcttttta atagcactaa ccaatgcctt tttagatgta
248
                                                                           1620
249
     tttttgatgt atatatctat tattcaaaaa atcatgttta ttttgagtcc taggacttaa
                                                                           1680
250 aattagtett ttgtaatate aageaggace etaagatgaa getgagettt tgatgeeagg
                                                                           1740
251 tgcaatctac tggaaatgta gcacttacgt aaaacatttg tttcccccac agttttaata
                                                                           1800
252 agaacagatc aggaattcta aataaatttc ccagttaaag attattgtga cttcactgta
                                                                           1860
253 tataaacata tttttatact ttattgaaag gggacacctg tacattcttc catcatcact
                                                                          1920
254 gtaaagacaa ataaatgatt atattcacaa aaaaaaaaa aaaaaa
                                                                          1966
256 <210> SEQ ID NO: 6
257 <211> LENGTH: 414
258 <212> TYPE: PRT
259 <213> ORGANISM: Unknown
261 <220> FEATURE:
262 <223> OTHER INFORMATION: predicted amino acid sequence of mouse dMTase1
264 <400> SEQUENCE: 6
265
    Met Arg Ala His Pro Gly Gly Gly Arg Cys Cys Pro Glu Gln Glu Glu
266
267
    Gly Glu Ser Ala Ala Gly Gly Ser Gly Ala Gly Gly Asp Ser Ala Ile
268
                20
                                     25
    Glu Gln Gly Gln Gly Ser Ala Leu Ala Pro Ser Pro Val Ser Gly
```

VERIFICATION SUMMARY

PATENT APPLICATION: US/09/554,414B

DATE: 11/12/2001

TIME: 18:23:52

Input Set : A:\corrected Sequence listing.txt Output Set: N:\CRF3\11122001\1554414B.raw

L:15 M:270 C: Current Application Number differs, Replaced Current Application No

L:15 M:271 C: Current Filing Date differs, Replaced Current Filing Date